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 Δ -6 pathway

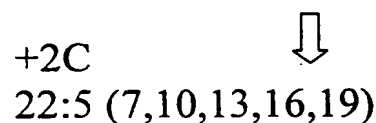
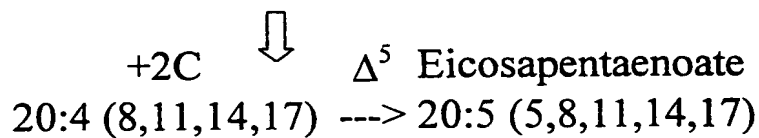
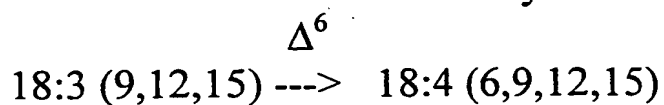
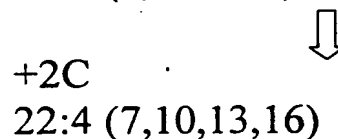
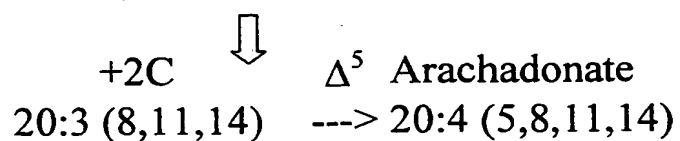
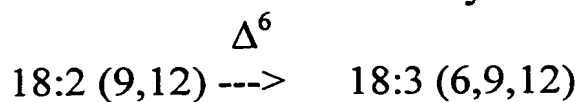
 ω -3 fatty acids ω -6 fatty acids

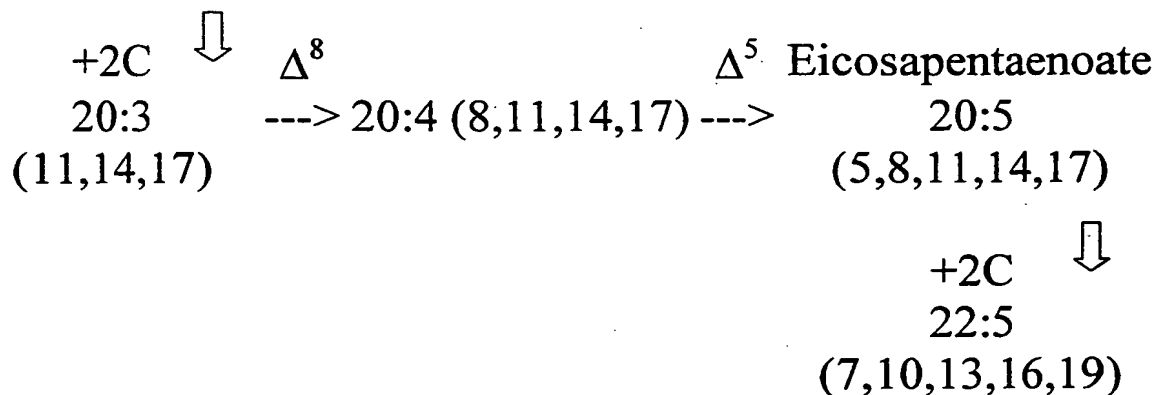
FIG. 1A

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Δ -8
pathway

ω -3 fatty acids

18:3 (9,12,15)



ω -fatty acids

18:2 (9,12)

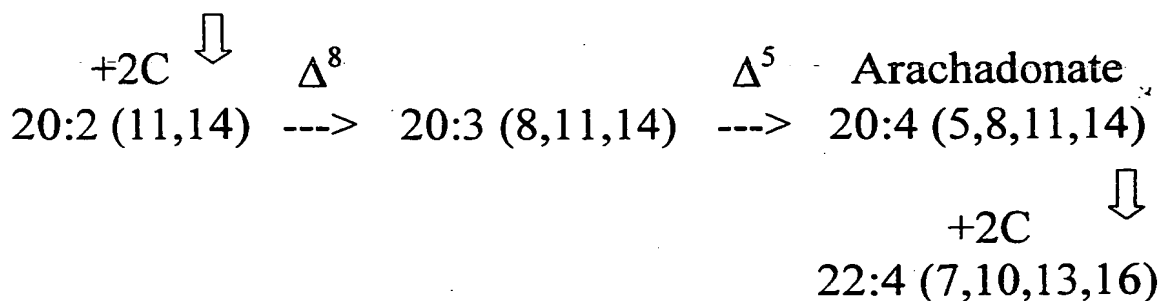


FIG. 1B

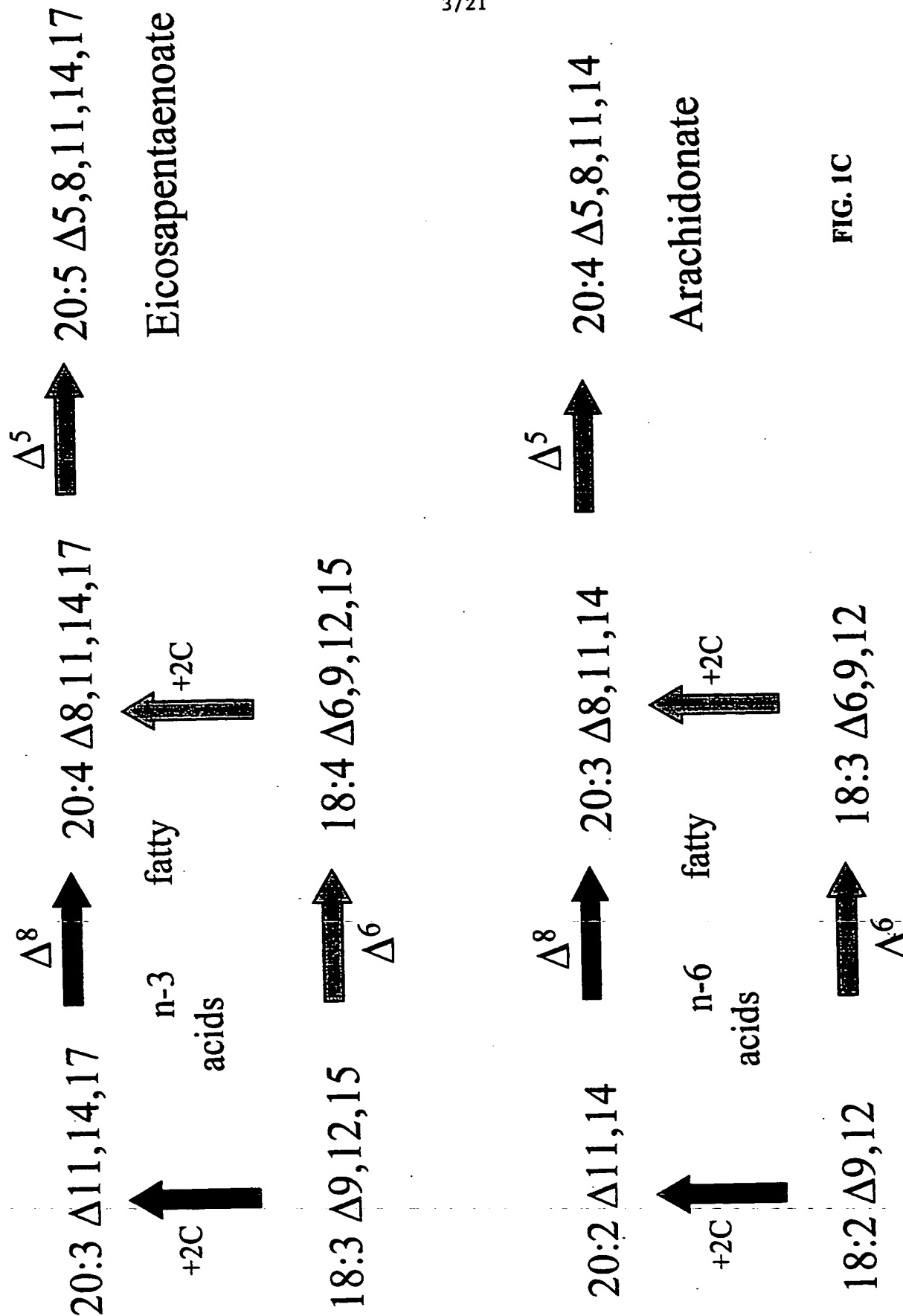
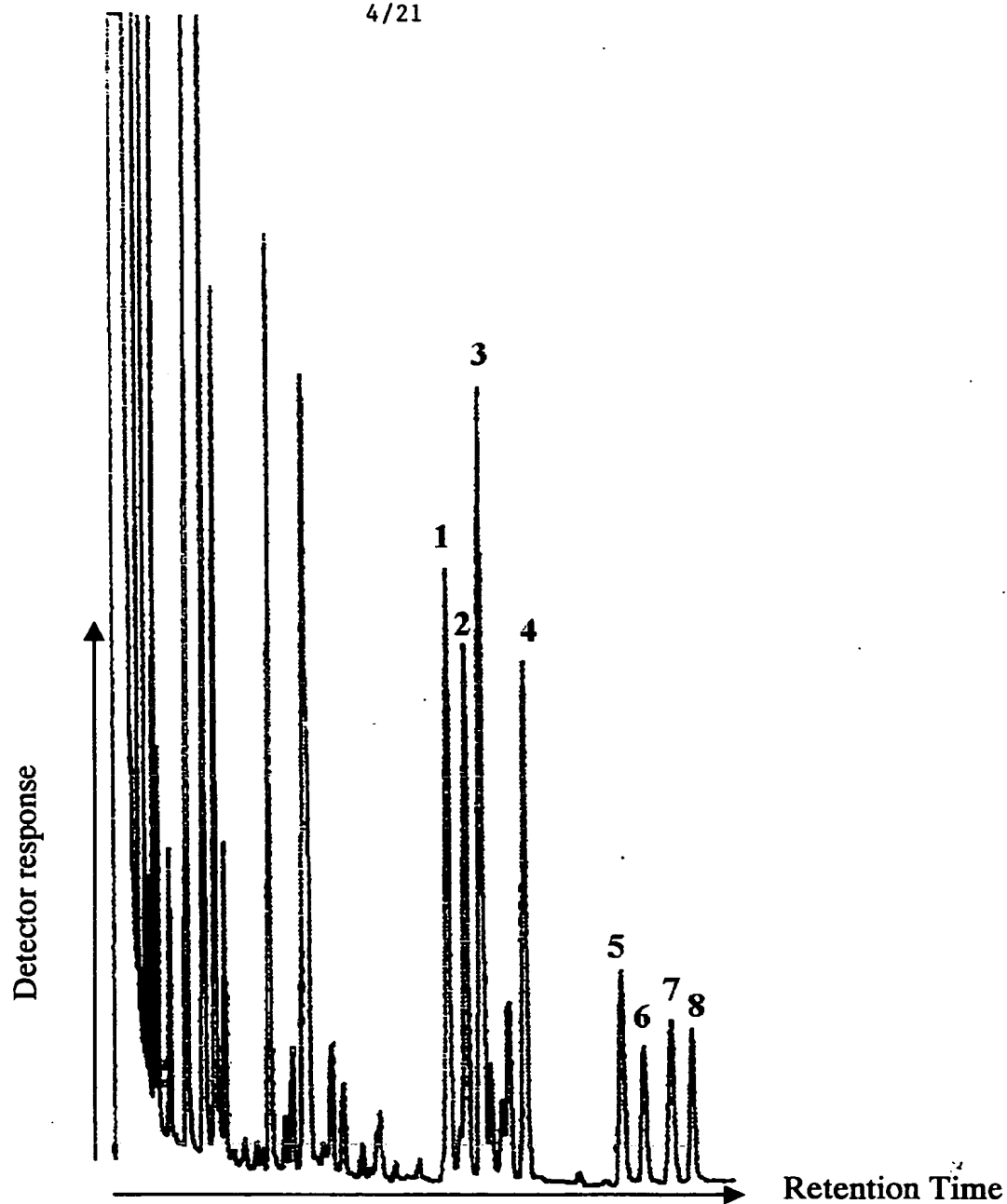


FIG. 1C

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PK	RT	FA	%	PK	RT	FA	
1	10.0	20:2 Δ 11,14	7.2	4	11.7	20:5 Δ 5,8,11,14,17	6.2
2	10.3	20:3 Δ 8,11,14	6.3	5	14.0	22:4 Δ 7,10,13,16	2.9
3	10.7	20:4 Δ 5,8,11,14	9.0	6	14.5	22:5 Δ 4,7,10,13,16	1.8
	10.9	20:3 Δ 11,14,17	1.4	7	15.2	22:5 Δ 7,10,13,16,19	2.6
				8	15.7	22:6 Δ 4,7,10,13,16,19	2.0

FIG. 2

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fat-4 1 MVLREQEHEPFFIKIDGKWCQIDDAVLRSHPGGS.AITTYKNMDATYVHTFTHTGSKRAY
 egd1 1 MKSKRQALSPLOEM...EQTYDVSAWVNEHPGGAETIENYQGRDATDAEMVMH...FOEAF
 fat-3 1 MVV.DKNASGLRMKVDGKWLYLSEELVKKHPGGA.VIEQYRNSDATHIFHAFHEGSSQAY

W HPGG G F H
 Cytochrome b₅-like domain

fat-4 60 QWLTELKKECPTQEPETIPDIKDDPIKGIDDVNMGTENISEKRSAGINKSETDLRMVRAE
 egd1 56 DKLRMPK.....INSEFELPPQAAVNEAQEDERKLREELIAT
 fat-3 59 KOLDLLKKH..GEHDEFLEKQLEKRLDKVDINVSAYDVSAQEKQVESEKLRKRLHOD

fat-4 120 GLMDGSPLEYIRKILETIFTILEFAFYLO.YHTYYLPSAILMGVAWQQLGWLITHEFAHHOL
 egd1 94 GMEDASPLWYSYKISTITLGLGVLYELMVQYQMYFICAVLLGMFYQMGWLSDICHHQT
 fat-3 117 GLMKANETYFLFKAISTLSIMAFAYLO.YLGWYITSACLLALAWQQLGWLITHEFCHQOP
HXKXHH

fat-4 179 FKNRYYNDLASYFVGNFLOGFSGCGWKEQHNVHHAATNVVGRDGLDLVFPFYATVAEHLN
 egd1 154 FKNRNWNNLVGLVFCNCLQCFSVTCWKDRHNAHHSATNVQGHDEPIDNLPPLAWSEDDVT
 fat-3 176 TKNRPLNDTISLFFGNFLOGFSRDWKKDKHNTTHHAATNVIDHDGIDLAPLFAFIPGDLG
HXKXHH

fat-4 239 NY...SODSWMTLFRWQHVVHTFMLELRLSWLLQSIIEVSOPTHYDYRNTAIYEQV
 egd1 214 RASPISRK...LIEFOQYYELVLCILLRFTWCFCVLTVRSIKDRDNOFYRSQYKKEAI
 fat-3 236 KYKASFEEKAILKIVPYOHLYFTAMLPLRFSWTCOSVQWVEKENOMEYKVYORNAFWEOA

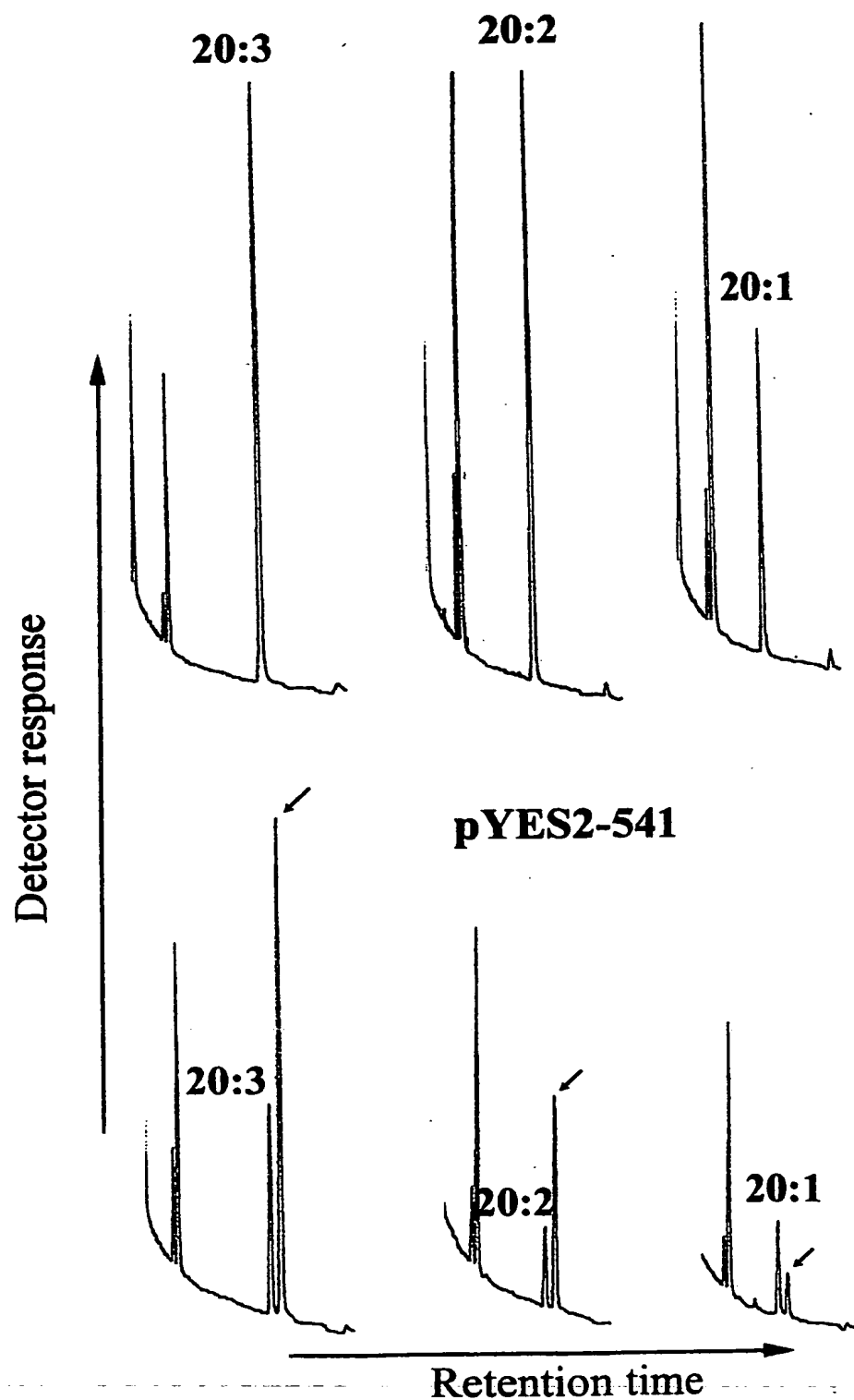
fat-4 297 GLSLHWAW.SLGQLYFLPDWSTRIMEFLVSHLVGGFLLSHVVTFNHYSVEKFALSSNIMS
 egd1 270 GLALHWTLKALEHLFFMPSTILTSLIVSEVSELVGGFCIATVVFVFNHYPLEKICDPVWDGH
 fat-3 296 THVGHAW.VFYQLFLLPTWPLRVAYETISQGGGLIAHVVTFNHNSVDKYPANSRTIN

fat-4 356 NYACLQIMTTRNMRPGREFIDWLWGGLNYQIEHHLFPTMPRHNLNTVMPLVKEFAAANGLP
 egd1 330 GFSVGQIHETMNIIRRCIITDWEFGGLNYQIEHHLWPTLPRHNLTAVSQOVEQLCQKHNL
 fat-3 355 NFAALQILTRNMTSPFIDWLWGGLNYQIEHHLFPTMPRONLNAOMKYVKEWCKENNL
HXKXHH

fat-4 416 YMVDDYFTGEWLEIEQERNIANVAAKITK.KIA
 egd1 390 YRNPLPHEGLVILLRYLAVEARMAEKOPACKAL
 fat-3 415 YLVDDYFDGYAMNLQOLKNMAE...HIOA.KAA

FIG. 3

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pYES2**FIG. 4**

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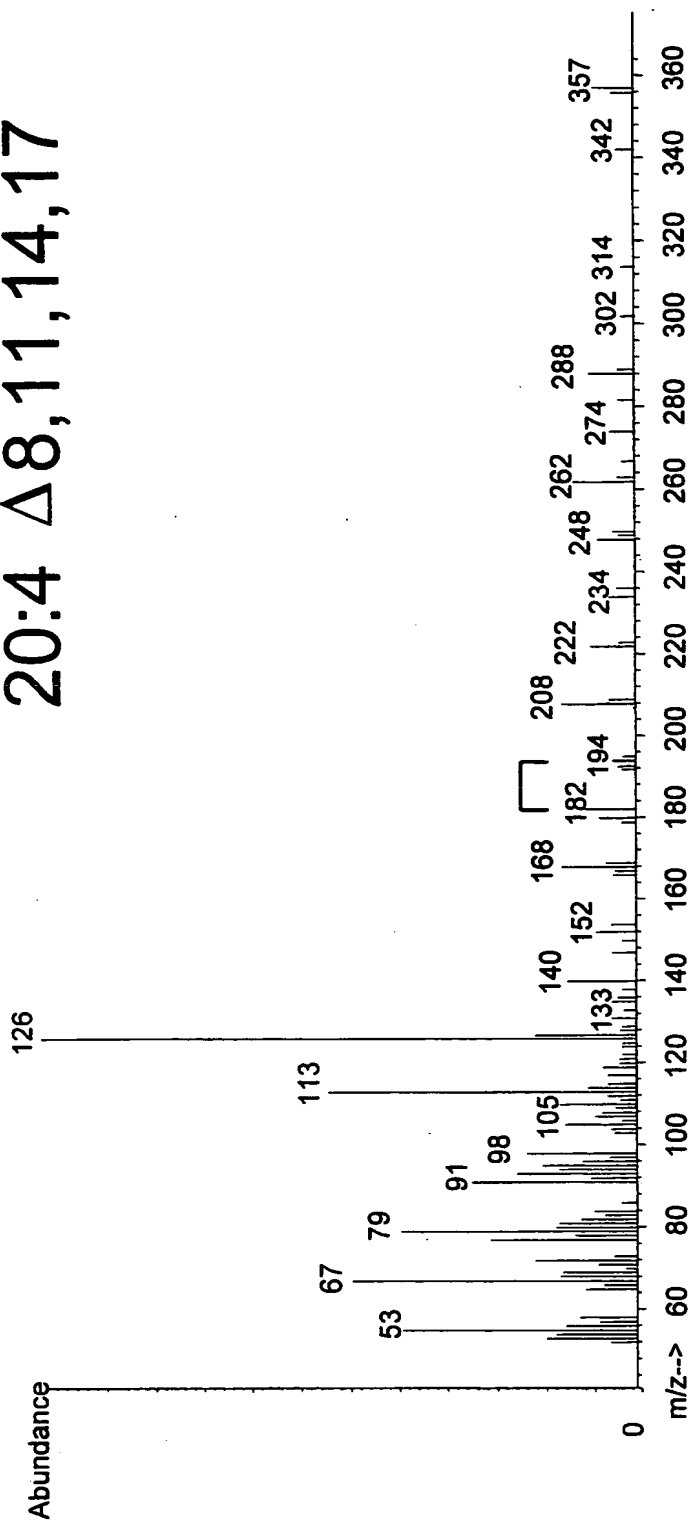
20:4 Δ 8,11,14,17

FIG. 5A

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20:3 Δ 8,11,14

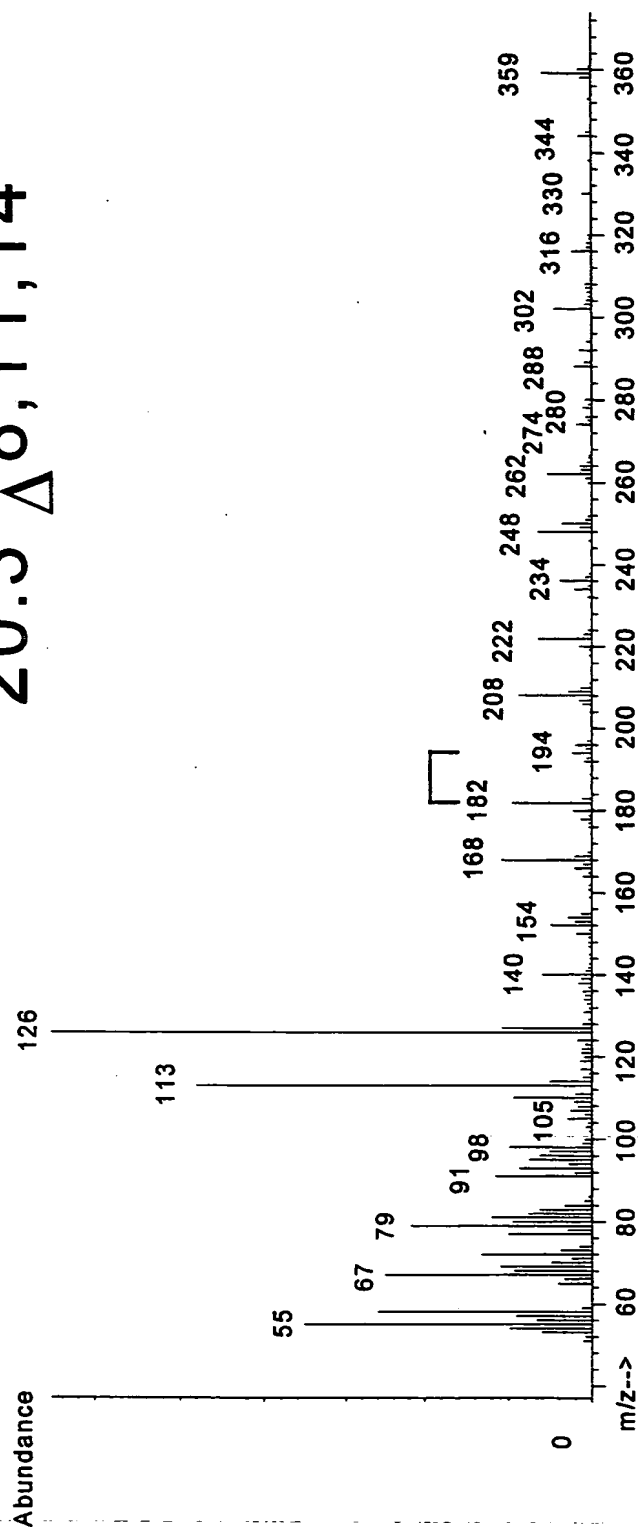


FIG. 5B

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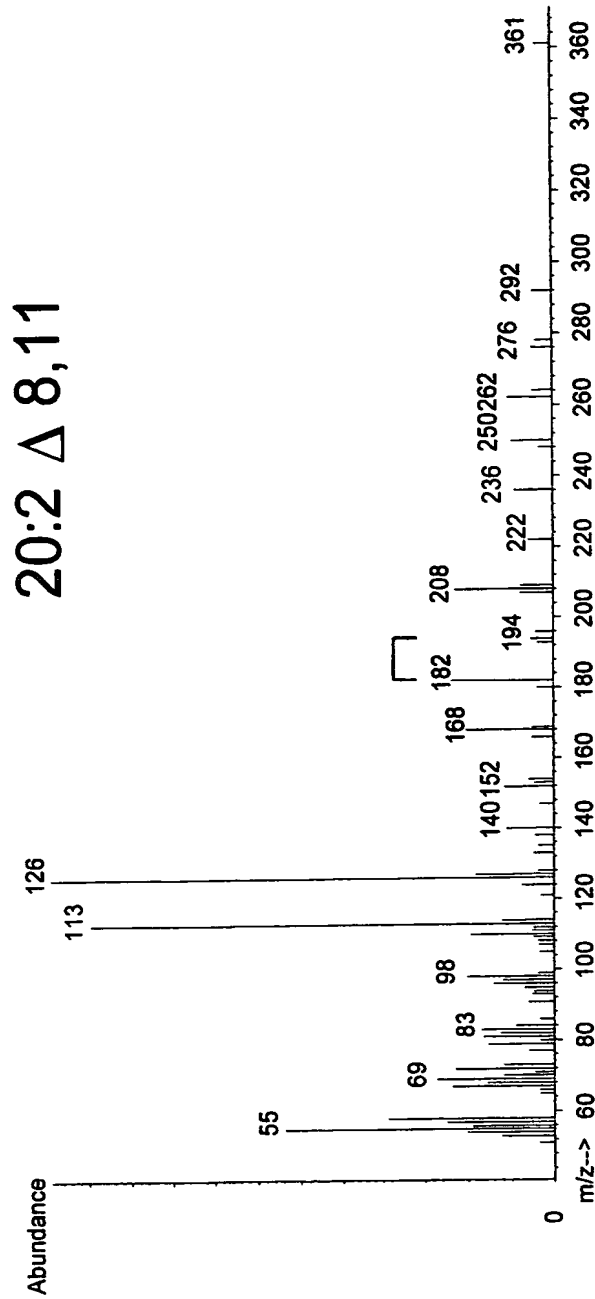


FIG. 5C

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1 MVLREQEHEP FFIKIDGKWC QIDDAVLRSH PGGSAITTYK NMDATTVFHT
51 FHTGSKEAYQ WLTELKKECP TQEPEIPDIK DDPIKGIDDV NMGTFNISEK
101 RSAQINKSFT DLRMVRRAEG LMDGSPLFYI RKILETIFTI LFAFYLQYHT
151 YYLPSAILMG VAWQQLGWL I HEFAHHQLFK NRYYN DLASY FVGNFLQGFS
201 SGGWKEQHN V HHAATNVVGR DGDLDLVPFY ATVAEHLNNY SQDSWVMTLF
251 RWQHVHWTFM LPFLRLSWLL QSIIFVSQMP THYYDYRNT AIYEQVGLSL
301 HWAWSLGQLY FLPDWSTKIM FFLVSHLVGG FLLSHVVTFN HYSVEKFALS
351 SNIMSNYACL QIMTTRNM RP GRFIDWLWGG LNYQIEHHLF PTMPRHNLT
401 VMPLVKEFAA ANGLPYMVDD YFTGFWLEIE QFRNIANVAA KLTKKIA

FIG. 6A

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```

1  GAATTTTCAA TCCTCCTTGG GTCCACCGC TGTGATATCA AAATGGTATT
51  ACGAGAGCAA GAGCATGAGC CATTCTTCAT TAAAAATTGAT GGAAAAATGGT
101  GTCAAATTGA CGATGCTGTC CTGAGATCAC ATCCAGGTGG TAGTGCAATT
151  ACTACCTATA AAAATATGGA TGCCACTACC GTATTCCACA CATTCCATAC
201  TGGTTCTAAA GAAGCGTATC AATGGCTGAC AGAATTGAAA AAAGAGTGCC
251  CTACACAAGA ACCAGAGATC CCAGATATTA AGGATGACCC AATCAAAGGA
301  ATTGATGATG TGAACATGGG AACTTTCAAT ATTTCTGAGA AACGATCTGC
351  CCAAATAAAT AAAAGTTTCA CTGATCTACG TATGCGAGTT CGTGCGAAG
401  GACTTATGGA TGGATCTCCT TTGTTCTACA TTAGAAAAAT TCTTGAAACA
451  ATCTTCACAA TTCTTTTTCG ATTCTACCTT CAATACCACA CATATTATCT
501  TCCATCAGCT ATTCTAATGG GAGTTGCGTG GCAACAATTG GGATGGTTAA
551  TCCATGAATT CGCACATCAT CAGTTGTTCA AAAACAGATA CTACAATGAT
601  TTGGCCAGCT ATTTCTGTTG AACTTTTTTA CAAGGATTCT CATCTGGTGG
651  TTGGAAAGAG CAGCACAATG TGCATCACGC AGCCACAAAT GTTGTGGAC
701  GAGACGGAGA TCTTGATTTA GTCCCATTCT ATGCTACAGT GGCAGAACAT
751  CTCAACAATT ATTCTCAGGA TTCATGGGTT ATGACTCTAT TCAGATGGCA
801  ACATGTTTCAT TGGACATTCA TGTTACCATT CCTCCGTCTC TCGTGGCTTC
851  TTCAGTCAAT CATTTTTGTT AGTCAGATGC CAACTCATT TATGACTAT
901  TACAGAAATA CTGCGATTTA TGAACAGGTT GGTCTCTCTT TGCACTGGGC
951  TTGGTCATTG GGTCaATTGT ATTTCTTACC CGATTGGTCA ACTAAAATAA
1001  TGTTCTTCCT TGTTTCTCAT CTTGTTGGAG GTTCTCTGCT CTCTCATGTA
1051  GTTACTTTCA ATCATTATTC AGTGGAGAAG TTTGCATTGA GCTCGAACAT
1101  CATGTCAAAT TACGCTTGTC TTCAAATCAT GACCACAAGA AATATGAGAC
1151  CTGGAAGATT CATTGACTGG CTTTGGGGAG GTCTTAACTA TCAGATTGAG
1201  CACCATCTTT TCCCAACGAT GCCACGACAC AACTTGAACA CTGTTATGCC
1251  ACTTGTTAAG GAGTTTGCAG CAGCAAATGG TTTACCATAC ATGGTCGACG
1301  ATTATTTTAC AGGATTCTGG CTTGAAATTG AGCAATTCCG AAATATTGCA
1351  AATGTTGCTG CTAaATTGAC TAAAAAGATT GCCTAGATTA CGATTAAATTA
1401  ATCAATTTAT TTTCATGTTT TATTCGTGTG TTTTAATATT TTCCAAATTT
1451  TTACCTATTC C

```

FIG. 6B

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1 MKSKRQALSP LQLMEQTYDV SAWVNFHPGG AEIIENYQGR DATDAFMVME
51 FQEAFDKLKR MPKINPSFEL PPQAAVNEAQ EDFRKLREEL IATGMFDASP
101 LWYSYKISTT LGLGVLGYFL MVQYQMYFIG AVLLGMHYQQ MGWLSHDICH
151 HQTfKNRnWN NLVGLVFGNG LQGFSVTCWK DRHNAHHSAT NVQGHDPDID
201 NLPPLAWSED DVTRASPISR KLIQFQQYYF LVICILLRFI WCFQCVLTVR
251 SLKDRDNQFY RSQYKKEAIG LALHWTLKAL FHLFFMPSIL TSLLVFFVSE
301 LVGGFGIAIV VFMNHYPLEK IGDVWDGHG FSVGQIHETM NIRRGITDW
351 FFGGLNYQIE HHLWPTLPRH NLTAVSYQVE QLCQKHNLPY RNPLPHEGLV
401 ILLRYLAVFA RMAEKQPAGK AL

FIG. 7A

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1 ATTTTTTTTC GAAATGAAGT CAAAGCGCCA AGCGCTATCC CCCTTACAAT
51 TGATGGAACA AACATATGAT GTGGTCAATT TCCACCCTGG TGGTGCGGAA
101 ATTATAGAGA ATTACCAAGG AAGGGATGCC ACTGATGCCT TCATGGTTAT
151 GCACTTTCAA GAAGCCTTCG ACAAGCTCAA GCGCATGCCC AAAATCAATC
201 CCAGTTTGA GTTGCCACCC CAGGCTGCAG TGAATGAAGC TCAAGAGGAT
251 TTCCGGAAGC TCCGAGAAGA GTTGATCGCA ACTGGCATGT TTGATGCCTC
301 CCCCCTCTGG TACTCATACA AAATCAGCAC CACACTGGGC CTTGGAGTGC
351 TGGGTTATTT CCTGATGGTT CAGTATCAGA TGTATTTTCAT TGGGGCAGTG
401 TTGCTTGGGA TGCACTATCA ACAGATGGGC TGGCTTTCTC ATGACATTTG
451 CCACCACCAG ACTTTCAAGA ACCGGAAGTGA GAACAACCTC GTGGGACTGG
501 TATTTGGCAA TGGTCTGCAA GGT'TTTTCCG TGACATGTTG GAAGGACAGA
551 CACAATGCAC ATCATTCGGC AACCAATGTT CAAGGGCAGC ACCCTGATAT
601 TGACAACTC CCCCCCTTAG CCTGGTCTGA GGATGACGTC ACACGGGCGT
651 CACCGATTC CCGCAAGCTC ATTCAGTTCC AGCAGTACTA TTTCTTGCTC
701 ATCTGTATCT TGTGCGGTT CATTTGGTGT TTCCAGTGCG TGTGACCGT
751 GCGCAGTTTG AAGGACAGAG ATAACCAATT CTATCGCTCT CAGTATAAGA
801 AGGAGGCCAT TGGCCTCGCC CTGCACTGGA CCTTGAAGGC CCTGTTCCAC
851 TTATTCTTTA TGCCCAGCAT CCTCACATCG CTGTTGGTGT TTTTCGTTTC
901 GGAGCTGGTT GCGGCTTCG GCATTGCGAT CGTGGTGTTC ATGAACCACT
951 ACCCACTGGA GAAGATCGGG GACCCAGTCT GGGATGGCCA TGGATTCTCG
1001 GTTGGCCAGA TCCATGAGAC CATGAACATT CGGCGAGGGA TTATCACAGA
1051 TTGTTTTC GGAGGCTTGA ATTACCAGAT TGAGCACCAT TTGTGGCCGA
1101 CCCTCCCTCG CCACAACCTG ACAGCGGTTA GCTACCAGGT GGAACAGCTG
1151 TGCCAGAAGC ACAACCTGCC GTATCGGAAC CCGCTGCCCC ATGAAGGGTT
1201 GGTCACTCTG CTGCGCTATC TGGCGGTGTT CGCCCGGATG GCGGAGAAGC
1251 AACCCGCGGG GAAGGCTCTA TAAGG

FIG. 7B

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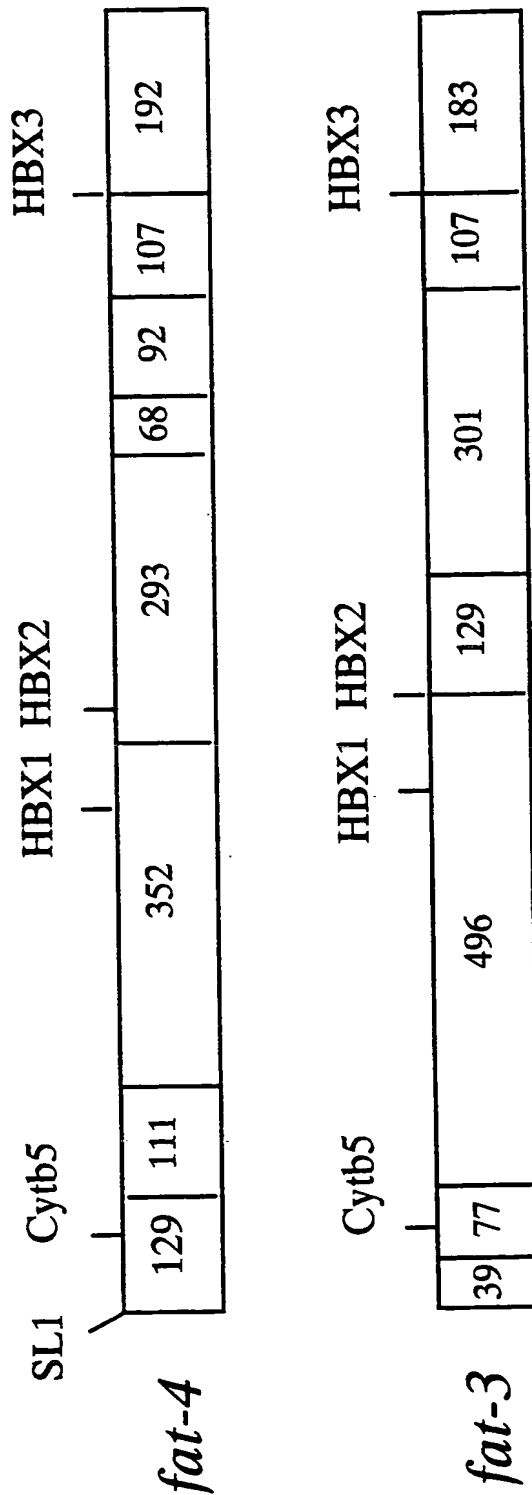
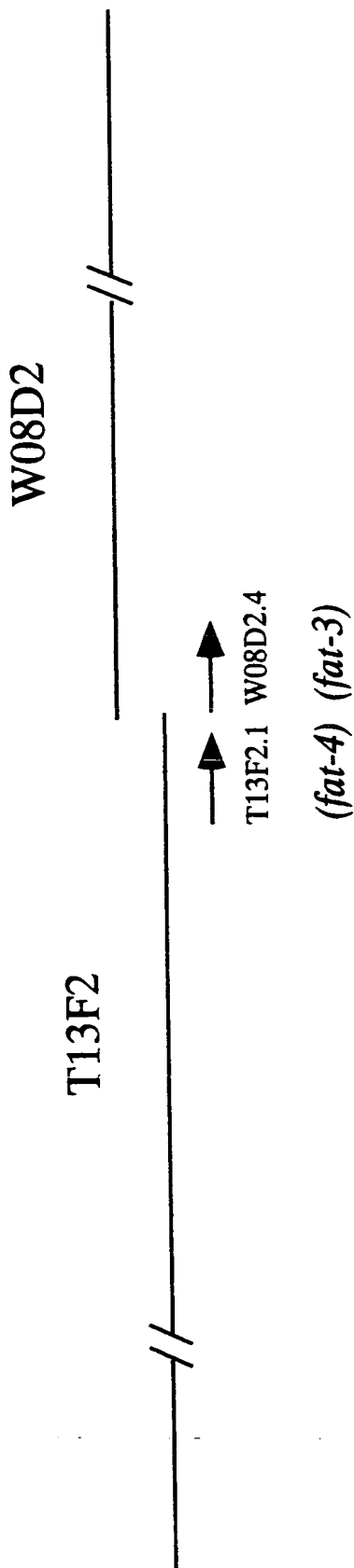


FIG. 8

FIG. 9

bord6	1	MMAQIKKY	ITSD	EMKNNHDK	KPGDL	IS	IQGKAY	DVSD	W	KRD	HPGGS	FPMK	LAGQ	MTDA
fat3	1	MV	AKNA	G	..	LRMK	VDCKWLYNS	EL	K	HPGGA	VIEQ	YRNS	DATH
fat4	1	MV	LR	GEHEP	..	FFIK	IDGKWC	DD	AV	SS	HPGGS	AIT	YKNMD
word5	1	M	QDQCKT	FT	WE	EA	HN	KGD	TE	FA	IT	GR	V	DV
bord6	60	F	VAFHPAS	..	T	K	LD	K	FT	C	Y	K	K	Y
fat3	47	F	HAFHE	GSSQ	AY	K	LD	L	K	K	..	G	E	H
fat4	48	F	HTEHT	GSKE	AY	Q	W	L	T	E	L	K	K	E
word5	60	E	EMH	A	G	A	D	A	M	K	K	Y	G
bord6	99	E	F	K	M	G	L	Y	D	R	K	G	H	E
fat3	105	S	F	E	K	L	R	O	Q	P	T	K	N	R
fat4	108	S	F	T	D	R	L	R	E	A	A	S	L	M
word5	106	Y	F	T	D	R	D	I	D	P	K	N	R	P
bord6	156	W	J	G	H	D	A	G	H	Y	M	V	S	D
fat3	165	W	L	T	H	E	F	C	H	O	Q	P	T	K
fat4	168	W	L	T	H	E	F	C	H	O	Q	P	T	K
word5	164	L	N	P	A	D	A	S	H	P	D	V	T	H
bord6	214	Y	P	F	L	V	S	K	E	F	G	S	T	S
fat3	223	L	A	P	L	E	A	P	L	E	A	P
fat4	226	L	E	P	L	E	A	P	L	E	A	P
word5	223	T	E	E	P	D	V	R	T	K	N	R	P	D
bord6	274	L	T	K	N	R	P	D	V	T	K	N	R	P
fat3	276	F	K	N	R	P	D	V	T	K	N	R	P	D
fat4	277	S	Q	M	P	T	H	Y	D	E	R	N	T	A
word5	279	P	I	S	T	W	H	A	V	M	A	G	C	K
bord6	327	F	S	L	N	H	E	S	S	V	Y	V	K	P
fat3	336	V	T	E	N	H	N	S	V	K	Y	P	K	N
fat4	337	V	T	E	N	H	N	S	V	K	Y	P	K	N
word5	339	V	Q	W	P	L	P	D	E	N	G	I	Q
bord6	385	C	N	L	R	K	S	P	Y	V	I	E	L	C
fat3	395	C	N	L	R	K	S	P	Y	V	I	E	L	C
fat4	396	H	N	L	T	V	M	P	L	V	K	E	A	A
word5	394	H	H	T	P	N	V	S	Q	H	H	Y	P
bord6	445	H	T	H	G								
fat3													
fat4													
word5	446	P	K	E	E								

FIG. 9

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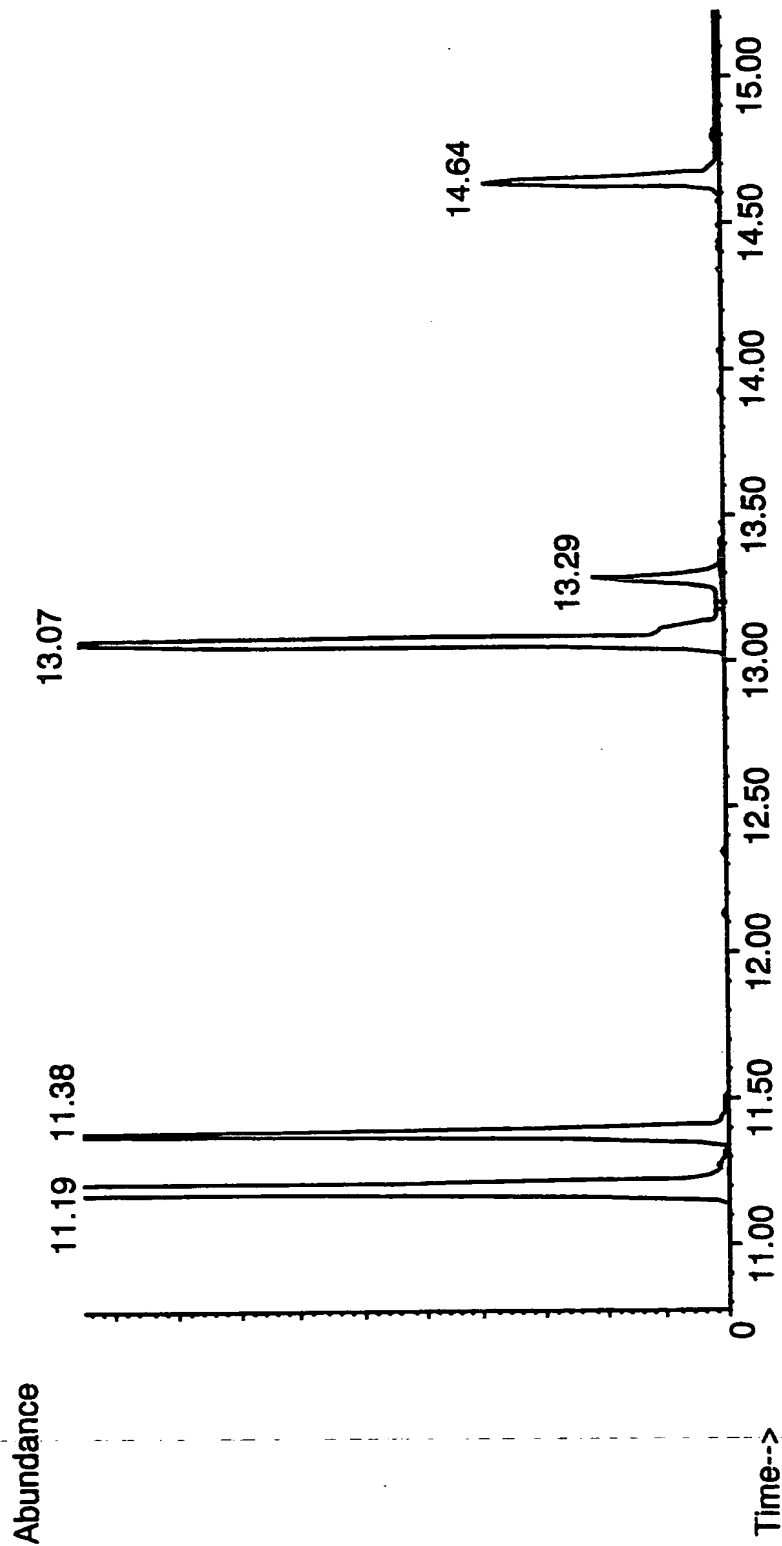


FIG. 10A

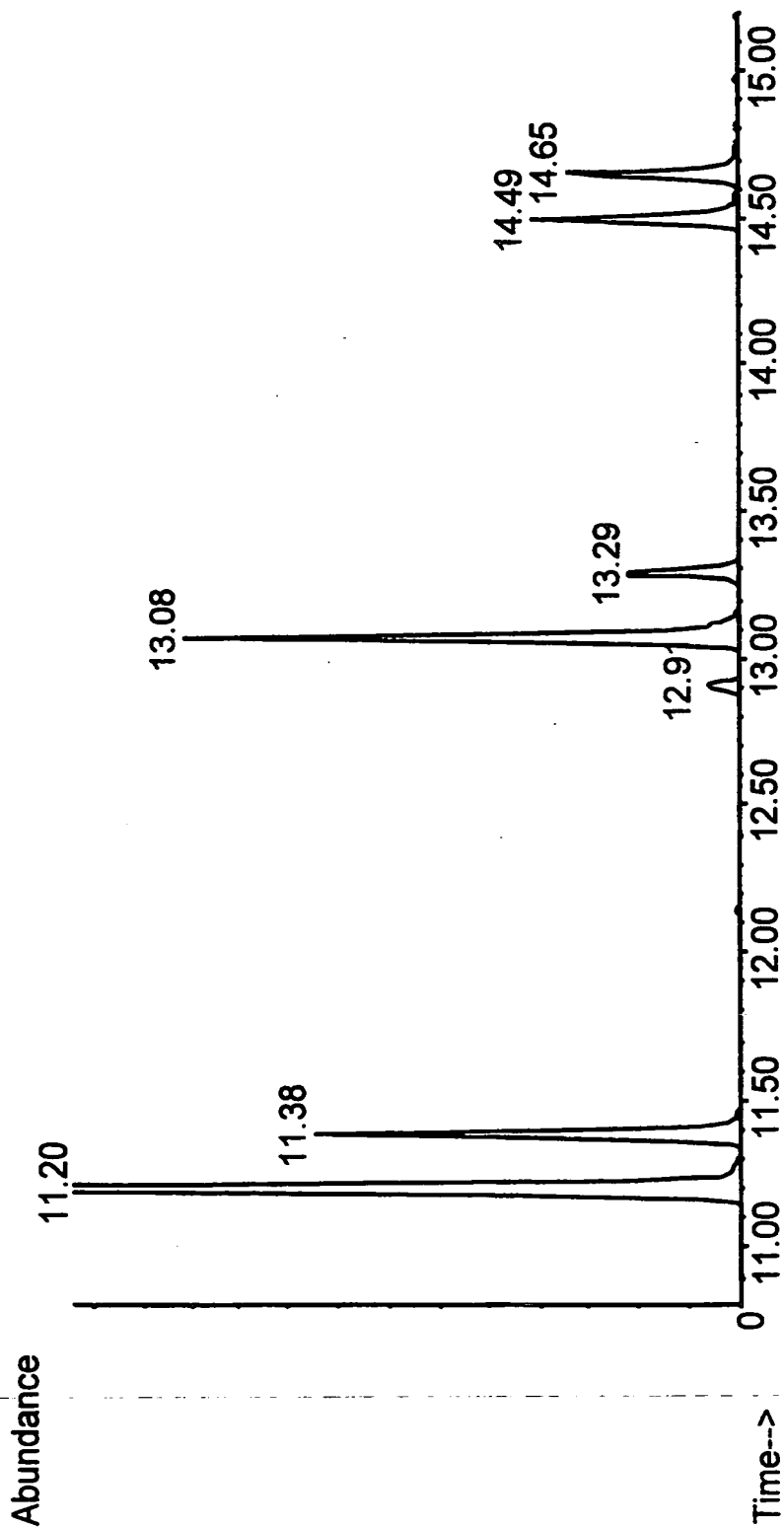


FIG. 10B

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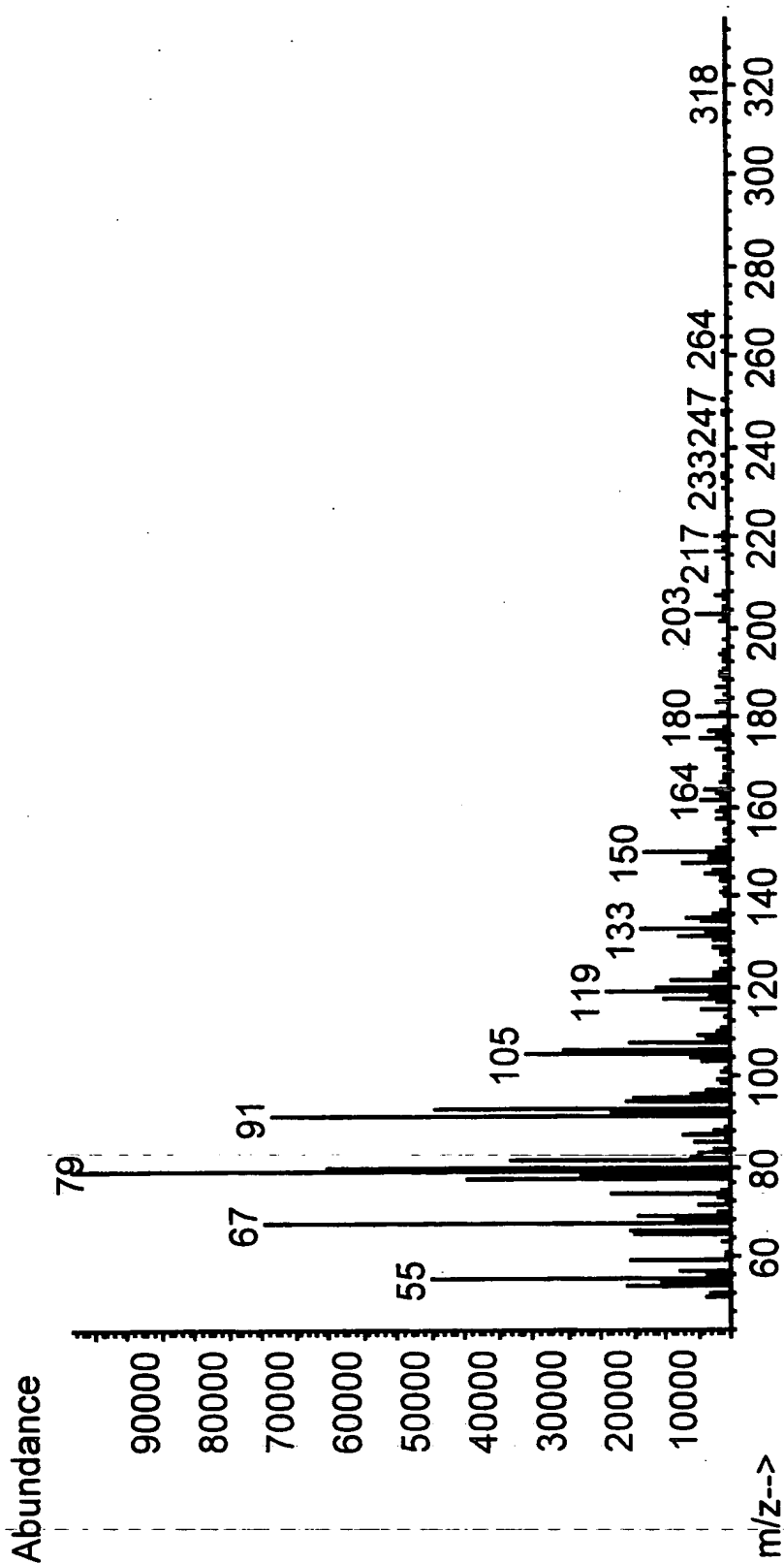
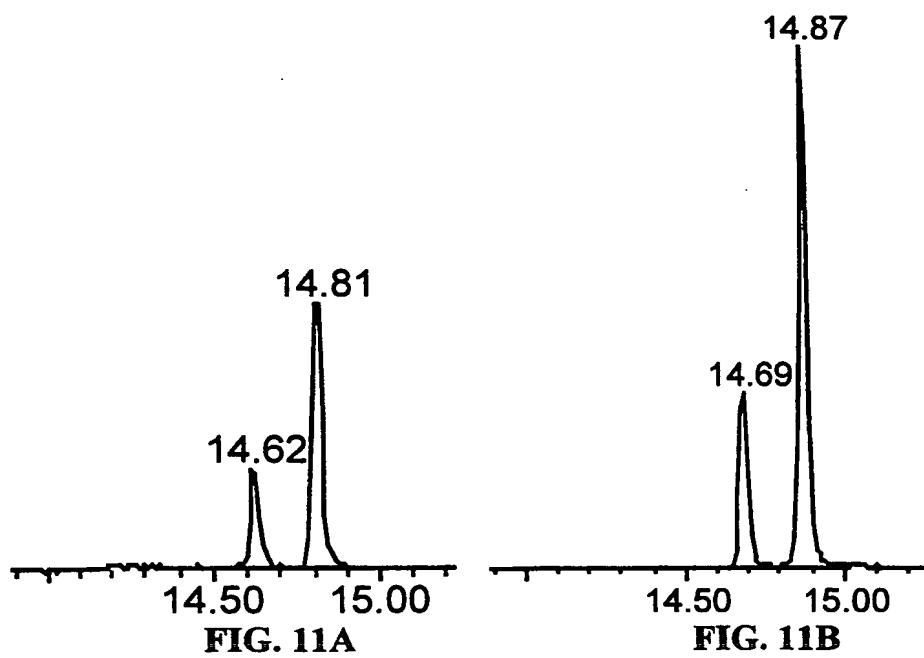


FIG. 10C

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FAT-3 (Δ^6 -desaturase)		FAT-4 (Δ^5 -desaturase)	
substrate	% of substrate converted	substrate	% of substrate converted
18:1 Δ^9	0	18:1 $\Delta^?$	5
18:2 $\Delta^9,12$	14	18:2 $\Delta^9,12$	0
18:3 $\Delta^9,12,15$	17	18:3 $\Delta^9,12,15$	0
20:1 Δ^{11}	0	20:1 Δ^{11}	0
20:2 $\Delta^{11,14}$	0	20:2 $\Delta^{11,14}$	27
20:3 $\Delta^{11,14,17}$	0	20:3 $\Delta^{11,14,17}$	26
20:3 $\Delta^8,11,14$	0	20:3 $\Delta^8,11,14$	55

FIG. 12

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	pYES		pYES-541	
	incorporation	desaturation	incorporation	desaturation
Δ^8 Substrates				
20:3 (11,14,17)	45.9	0	38.4	27.2
20:2 (11,14)	16.7	0	21.3	14.8
20:1 (11)	15.5	0	18.8	6.1
Δ^6 Substrates				
18:3 (9,12,15)	20.9	0	19.2	0
18:2 (9,12)	15.8	0	18.5	0
Δ^5 Substrate				
20:3 (8,11,14)	34.4	0	35.6	0

FIG. 13

09/857583-001701